## PATENT COOPERATION TREATY

# PCT

#### INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference 01107.00113	FOR FURTHER see Notification of Transmittal of International Search Report (Form PCT/ISA/220) as well as, where applicable, item 5 below.		
International application No.	International filing date (day/month/year)	(Earliest) Priority Date (day/month/year)	
PCT/US 01/08886	21/03/2001	21/03/2000	
Applicant THE JOHNS UNIVERSITY et a	1.		
This International Search Report has bee according to Article 18. A copy is being tra	n prepared by this International Searching Aut ansmitted to the International Bureau.	thority and is transmitted to the applicant	
This International Search Report consists  It is also accompanied by	of a total of <u>6</u> sheets.  a copy of each prior art document cited in this	s report.	
Basis of the report			
	international search was carried out on the ba less otherwise indicated under this item.	ssis of the international application in the	
the international search w Authority (Rule 23.1(b)).	vas carried out on the basis of a translation of	the international application furnished to this	
<ul> <li>With regard to any nucleotide an was carried out on the basis of the</li> </ul>	•	nternational application, the international search	
	onal application in written form.		
filed together with the inte	ernational application in computer readable for	m.	
T furnished subsequently to	this Authority in written form.		
furnished subsequently to	this Authority in computer readble form.		
	osequently furnished written sequence listing one is filed has been furnished.	does not go beyond the disclosure in the	
the statement that the info furnished	ormation recorded in computer readable form i	is identical to the written sequence listing has been	
2. X Certain claims were fou	nd unsearchable (See Box I).		
3. Unity of invention is lac	king (see Box II).		
4. With regard to the title,			
X the text is approved as su	bmitted by the applicant.		
the text has been establis	hed by this Authority to read as follows:		
		ity as it appears in Box III. The applicant may, port, submit comments to this Authority.	
6. The figure of the <b>drawings</b> to be publ	ished with the abstract is Figure No.	1a	
as suggested by the appli	cant.	None of the figures.	
$oxed{X}$ because the applicant fail	ed to suggest a figure.		
because this figure better	characterizes the invention.		

A. CLASSI IPC 7	IFICATION OF SUBJECT MATTER CO7K7/08 A61K47/48 A61K38/	10	
			* 4 * 3
	o International Patent Classification (IPC) or to both national classific	ation and IPC	
	SEARCHED ocumentation searched (classification system followed by classification)	ion symbols)	
IPC 7	CO7K A61K		
Documenta	tion searched other than minimum documentation to the extent that s	such documents are included in the fields so	earched
Electronic d	data base consulted during the international search (name of data ba	ase and, where practical, search terms used	1)
EPO-In	ternal, WPI Data, PAJ, CHEM ABS Data	a, MEDLINE, BIOSIS	
C. DOCUM	ENTS CONSIDERED TO BE RELEVANT		
Category °	Citation of document, with indication, where appropriate, of the re	levant passages	Relevant to claim No.
Y	RICOUART A ET AL: "DESIGN OF POPROTEIN KINASE INHIBITORS USING BISUBSTRATE APPROACH" JOURNAL OF MEDICINAL CHEMISTRY, CHEMICAL SOCIETY. WASHINGTON, US vol. 34, no. 1, 1991, pages 73-78 XP002918324 ISSN: 0022-2623 See especially page 73, column 1 column 1	THE AMERICAN , B,	1-68
X Furti	her documents are listed in the continuation of box C.	Patent family members are listed	in annex.
° Special ca	ategories of cited documents :		
<ul> <li>"A" document defining the general state of the art which is not considered to be of particular relevance</li> <li>"E" earlier document but published on or after the international filing date</li> <li>"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)</li> <li>"O" document referring to an oral disclosure, use, exhibition or other means</li> <li>"P" document published prior to the international filing date but</li> </ul>		<ul> <li>"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention</li> <li>"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone</li> <li>"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.</li> </ul>	
		*&" document member of the same patent Date of mailing of the international sea	Tarriny
Date of the actual completion of the international search  5. February 2002		27/02/2002	агон Героп
Name and mailing address of the ISA		Authorized officer	
European Palent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016		Groenendijk, M	

PCT/US 01/08886

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT  Relevant to claim No.				
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Helevani to claim No.		
Y .	HUBBARD S R: "Crystal structure of the acfivated insulin receptor tyrosine kinase in complex with peptide substrate and ATP analog" EMBO JOURNAL, OXFORD UNIVERSITY PRESS, SURREY, GB, vol. 16, no. 18, 1997, pages 5572-5581, XP002088043 ISSN: 0261-4189 cited in the application The whole document; see especially page 5578, column 2	1-68		
Ρ,Χ	MILLER: "Double trouble"  NATURE STRUCTURAL BIOLOGY,  vol. 8, no. 1, January 2001 (2001-01),  pages 16-18, XP001053333  the whole document	1-68		
P,X	PARANG E.A.: "Mechanism-based design of a protein kinase inhibitor" NATURE STRUCTURAL BIOLOGY, vol. 8, no. 1, January 2001 (2001-01), pages 37-41, XP001053332 the whole document	1-68		
Y	MEDZIHRADSZKY DENES ET AL: "Solid-phase synthesis of adenosine phosphopeptides as potential bisubstrate inhibitors of protein kinases"  JOURNAL OF THE AMERICAN CHEMICAL SOCIETY, AMERICAN CHEMICAL SOCIETY, WASHINGTON, DC, US, vol. 116, no. 21, 1994, pages 9413-9419, XP002150234 ISSN: 0002-7863 cited in the application the whole document	30-57, 59-68		
A	ROSSE, GERARD ET AL: "Synthesis of modified tripeptides and tetrapeptides as potential bisubstrate inhibitors of the epidermal growth factor receptor protein tyrosine kinase" HELV. CHIM. ACTA (1997), 80(3), 653-670, XP002188862 See especially page 654  -/	1-68		

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International Application No PCT/US 01/08886

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT Relevant to claim No. Citation of document, with indication, where appropriate, of the relevant passages 1-68 "Kinetic analysis of a protein Α KIM E.A.: tyrosine kinase reaction transition state in the forward and reverse directions" JOURNAL OF THE AMERICAN CHEMICAL SOCIETY, vol. 120, no. 28, 22 July 1998 (1998-07-22), pages 6851-6858, XP002188863 DC US the whole document

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#### FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.2

Claims Nos.: 1-4,8-19,23-33,37-47,51-68

Present claims 1,16,30,44 and 60 relate to an extremely large number of possible compounds (and their use). Support within the meaning of Article 6 PCT and/or disclosure within the meaning of Article 5 PCT is to be found, however, for only a very small proportion of the compounds claimed. In the present case, the claims so lack support, and the application so lacks disclosure, that a meaningful search over the whole of the claimed scope is impossible. Moreover the majority of said compounds is only defined by a minimum length of the tether, lacking any other structural properties. Hence said claims also lack clarity within the meaning of Article 6 PCT to such an extent as to render a meaningful search of the whole of the claimed scope of said claims impossible. Consequently, the search has been carried out for those parts of the claims which appear to be clear, supported and disclosed, namely those parts relating to the compounds defined in the claims 5-7 and 34-36 and their use and also for the general concept underlying the claims 1,16,30,44,and 60.

The applicant's attention is drawn to the fact that claims, or parts of claims, relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure.

International application No. PCT/US 01/08886

### INTERNATIONAL SEARCH REPORT

Box	Observations where certain claims were found unsearchable (Continuation of it m 1 of first sheet)
This Int	ernational Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X	Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
	Although claims 16-29 and 44-57 encompass or are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
2. X	Claims Nos.: 1–4,8–19,23–33,37–47,51–68 because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
	see FURTHER INFORMATION sheet PCT/ISA/210
	· ·
3	Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II	Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This Inte	ernational Searching Authority found multiple inventions in this international application, as follows:
	·
1.	As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2.	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3.	As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4.	No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
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Remark	The additional search fees were accompanied by the applicant's protest.
	No protest accompanied the payment of additional search fees.
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